

|   |   |                        |
|---|---|------------------------|
| <b>INFORMATION<br/>DISCLOSURE<br/>STATEMENT</b> | Atty. Docket No.: 6283.N DV1  | Serial No.: 10/694,385 |
|   | Applicant(s): Stockman et al.   | Confirmation No.: 5758 |
|   | Application Filing Date: Oct. 27, 2003  | Group: 1645            |
|   | Information Disclosure Statement<br>submitted (via facsimile) on: <b>MARCH</b> <u>  </u> / <u>  </u> , 2004 |                        |

**U.S. PATENT DOCUMENTS**

| Examiner<br>Initial | Copy<br>Enclosed | Document Number | Date     | Name             | Class | Subclass | Filing Date If<br>Appropriate |
|---------------------|------------------|-----------------|----------|------------------|-------|----------|-------------------------------|
| MS                  |                  | 4,719,582       | 01/12/88 | Ishida et al.    |       |          |                               |
|                     |                  | 5,270,163       | 12/14/93 | Gold et al.      |       |          |                               |
|                     |                  | 5,306,619       | 04/26/94 | Edwards et al.   |       |          |                               |
|                     |                  | 5,668,734       | 09/16/97 | Krishna et al.   |       |          |                               |
|                     |                  | 5,698,401       | 12/16/97 | Fesik et al.     |       |          |                               |
|                     |                  | 5,804,390       | 09/08/98 | Fesik et al.     |       |          |                               |
|                     |                  | 5,837,460       | 11/17/98 | Von Feldt et al. |       |          |                               |
|                     |                  | 5,856,496       | 01/05/99 | Fagnola et al.   |       |          |                               |
|                     |                  | 5,891,643       | 04/06/99 | Fesik et al.     |       |          |                               |
|                     |                  | 5,989,827       | 11/23/99 | Fesik et al.     |       |          |                               |
|                     |                  | 6,043,024       | 03/28/00 | Fesik et al.     |       |          |                               |
|                     |                  | 6,214,561       | 04/10/01 | Peters et al.    |       |          |                               |
|                     |                  | 6,677,160 B1    | 01/13/04 | Stockman et al.  |       |          |                               |
|                     |                  | 2001/0051333 A1 | 12/13/01 | Stockman         |       |          |                               |
| ↓                   |                  | 2002/0192701 A1 | 12/19/02 | Stockman et al.  |       |          |                               |

**FOREIGN PATENT DOCUMENTS**

| Examiner<br>Initial | Copy<br>Enclosed | Document Number        | Date     | Country                                     | Class | Subclass | Translation |    |
|---------------------|------------------|------------------------|----------|---|-------|----------|-------------|----|
|                     |                  |                        |          |   |       |          | Yes         | No |
| MS                  |                  | EP 0 592,816 A1,<br>B1 | 04/20/94 | EPO (with English<br>language abstract)     |       |          |             | X  |
|                     |                  | DE 196 49 359 C1       | 02/12/98 | Germany (with English<br>language abstract) |       |          |             | X  |
|                     |                  | GB 2 316 941 A         | 03/11/98 | Great Britain                               |       |          |             |    |
| ↓                   |                  | GB 2 321 104 A         | 07/15/98 | Great Britain                               |       |          |             |    |

|  |  |
|--|--|
| <b>EXAMINER</b><br><br>/Mark Shibuya/  | <b>Date Considered</b><br><br>06/21/2006 |
| *Examiner: Initial if citation considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant. |  |

|   |   |                        |
|---|---|------------------------|
| <b>INFORMATION<br/>DISCLOSURE<br/>STATEMENT</b> | Atty. Docket No.: 6283.N DV1  | Serial No.: 10/694,385 |
|   | Applicant(s): Stockman et al.   | Confirmation No.: 5758 |
|   | Application Filing Date: Oct. 27, 2003  | Group: 1645            |
|   | Information Disclosure Statement<br>submitted (via facsimile) on: <b>MARCH 1</b> , 2004 |                        |

|    |  |             |          |     |  |  |  |  |
|----|--|-------------|----------|-----|--|--|--|--|
| MS |  | WO 91/10140 | 07/11/91 | PCT |  |  |  |  |
|    |  | WO 91/17428 | 11/14/91 | PCT |  |  |  |  |
|    |  | WO 93/00446 | 01/07/93 | PCT |  |  |  |  |
|    |  | WO 94/14980 | 07/07/94 | PCT |  |  |  |  |
|    |  | WO 96/30849 | 10/03/96 | PCT |  |  |  |  |
|    |  | WO 97/00244 | 01/03/97 | PCT |  |  |  |  |
|    |  | WO 97/18469 | 05/22/97 | PCT |  |  |  |  |
|    |  | WO 97/18471 | 05/22/97 | PCT |  |  |  |  |
|    |  | WO 98/46548 | 10/22/98 | PCT |  |  |  |  |
|    |  | WO 98/48264 | 10/29/98 | PCT |  |  |  |  |
|    |  | WO 98/57155 | 12/17/98 | PCT |  |  |  |  |
|    |  | WO 99/09024 | 02/25/99 | PCT |  |  |  |  |
|    |  | WO 99/17616 | 04/15/99 | PCT |  |  |  |  |
|    |  | WO 99/36422 | 07/22/99 | PCT |  |  |  |  |
|    |  | WO 99/43643 | 09/02/99 | PCT |  |  |  |  |
|    |  |             |          |     |  |  |  |  |
|    |  |             |          |     |  |  |  |  |

**OTHER DOCUMENTS (Including Authors, Title, Date, Pertinent Papers, etc.)**

| Examiner Initial | Copy Enclosed | Document Description  |
|------------------|---------------|---|
| MS               |               | Ajay et al., "Can We Learn To Distinguish Between "Drug-like" and "Nondrug-like" Molecules?," <i>J. Med. Chem.</i> , 41:3314-3324 (1998).   |
|                  |               | Anderson et al., "Affinity NMR: Decoding DNA Binding," <i>Journal of Combinatorial Chemistry</i> , 1(1):69-72 (1999).   |
|                  |               | Balaram et al., "Localization of Tyrosine at the Binding Site of Neurophysin II by Negative Nuclear Overhauser Effects," <i>Journal of the American Chemical Society</i> , 94(11):4017-4018 (1972). |

|  |  |
|--|--|
| <b>EXAMINER</b><br><br>/Mark Shibuya/  | <b>Date Considered</b><br><br>06/21/2006 |
| *Examiner: Initial if citation considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant. |  |

|   |  |                               |
|---|--|-------------------------------|
| <b>INFORMATION<br/>DISCLOSURE<br/>STATEMENT</b> | <b>Atty. Docket No.:</b> 6283.N DV1  | <b>Serial No.:</b> 10/694,385 |
|   | <b>Applicant(s):</b> Stockman et al.   | <b>Confirmation No.:</b> 5758 |
|   | <b>Application Filing Date:</b> Oct. 27, 2003  | <b>Group:</b> 1645            |
|   | Information Disclosure Statement<br>submitted (via facsimile) on: <b>MARCH 1, 2004</b> |                               |

| Examiner<br>Initial | Copy<br>Enclosed | Document Description   |
|---------------------|------------------|--|
| MS                  |                  | Barjat et al., "High-Resolution Diffusion-Ordered 2D Spectroscopy (HR-DOSY) - A New Tool for the Analysis of Complex Mixtures," <i>Journal of Magnetic Resonance, Series B</i> , 108:170-172 (1995).                                   |
|                     |                  | Bax et al., "Sensitivity-Enhanced Two-Dimensional Heteronuclear Shift Correlation NMR Spectroscopy," <i>Journal of Magnetic Resonance</i> , 67:565-569 (1986).   |
|                     |                  | Belton et al., "Application of Chemometrics to the <sup>1</sup> H NMR Spectra of Apple Juices: Discrimination Between Apple Varieties," <i>Food Chemistry</i> , 61:207-213 (1998).   |
|                     |                  | Bemis et al., "The Properties of Known Drugs. 1. Molecular Frameworks," <i>J. Med. Chem.</i> , 39:2887-2893 (1996).  |
|                     |                  | Bemis et al., "Properties of Known Drugs. 2. Side Chains," <i>J. Med. Chem.</i> , 42:5095-5099 (1999).   |
|                     |                  | BLAST 2 Sequences. [online] National Center for Biotechnology Information, National Institutes of Health, United States, [retrieved 2001-08-29]. Retrieved from the Internet: <URL:http://www.ncbi.nlm.nih.gov/gorf/bl2.html>, 1 page. |
|                     |                  | Bleicher et al., "Diffusion Edited NMR: Screening Compound Mixtures by Affinity NMR to Detect Binding Ligands to Vancomycin," <i>Journal of Organic Chemistry</i> , 63(23):8486-8490 (1998).   |
|                     |                  | Boguslavsky, "NMR Finds Elusive Protein-Binding Molecules," <i>Drug Discovery &amp; Development</i> , 56-60 (1999).  |
| ✓                   |                  | Bothner-By et al., "Binding of Small Molecules to Proteins," <i>Ann. NY Acad. Sci.</i> , 222:668-675 (1973).   |
|                     |                  | <del>Braker <a href="http://www.braker.de/analytic/nmr-dep/best/best.htm">http://www.braker.de/analytic/nmr-dep/best/best.htm</a></del>  |
| MS                  |                  | Bussiere et al., "Structure of the E2 DNA-Binding Domain form Human Papillomavirus Serotype 31 at 2.4 Å," <i>Acta Crystallographica, D54</i> (Part 6, No. 2):1367-1376 (1998).   |
| MS                  |                  | Chen et al., "NOE Pumping: A Novel NMR Technique for Identification of Compounds with Binding Affinity to Macromolecules," <i>Journal of the American Chemical Society</i> , 120(39):10258-10259 (1998).                               |

|  |  |
|--|--|
| <b>EXAMINER</b><br><br>/Mark Shibuya/  | <b>Date Considered</b><br><br>06/21/2006 |
| *Examiner: Initial if citation considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant. |  |

|   |  |                               |
|---|--|-------------------------------|
| <b>INFORMATION<br/>DISCLOSURE<br/>STATEMENT</b> | <b>Atty. Docket No.:</b> 6283.N DV1  | <b>Serial No.:</b> 10/694,385 |
|   | <b>Applicant(s):</b> Stockman et al.   | <b>Confirmation No.:</b> 5758 |
|   | <b>Application Filing Date:</b> Oct. 27, 2003  | <b>Group:</b> 1645            |
|   | Information Disclosure Statement<br>submitted (via facsimile) on: <b>MARCH 1, 2004</b> |                               |

| Examiner<br>Initial | Copy<br>Enclosed | Document Description  |
|---------------------|------------------|---|
| MS                  |                  | Chen et al., "NOE Pumping. 2. A High-Throughput Method To Determine Compounds with Binding Affinity to Macromolecules by NMR," <i>J. Am. Chem. Soc.</i> , 122:414-415 (2000).   |
|                     |                  | Chiyoda et al., "Screening System for Urease Inhibitors Using <sup>13</sup> C-NMR," <i>Chemical &amp; Pharmaceutical Bulletin</i> , 46(4):718-720 (1998).   |
|                     |                  | Dalvit et al., "Sensitivity-Improved Detection of Protein Hydration and Its Extension to the Assignment of Fast-Exchanging Resonances," <i>J. Magn. Reson. B.</i> 109:334-338 (1995).   |
|                     |                  | Dalvit, "Homonuclear 1D and 2D NMR Experiments for the Observation of Solvent-Solute Interactions," <i>J Magn Reson B.</i> 1996 Sep;112(3):282-288.   |
|                     |                  | Dalvit et al., "Half-Filter Experiments for Assignment, Structure Determination and Hydration Analysis of Unlabelled Ligands Bound to <sup>13</sup> C/ <sup>15</sup> N Labelled Proteins," <i>J. Biomol. NMR</i> , 13:43-50 (1999). |
|                     |                  | Dalvit et al., "Use of Organic Solvents and Small Molecules for Locating Binding Sites on Proteins in Solution," <i>J. Biomol. NMR</i> , 14(1):23-32 (1999).  |
|                     |                  | Dalvit et al., "Identification of Compounds with Binding Affinity to Proteins Via Magnetization Transfer From Bulk Water," <i>Journal of Biomolecular NMR</i> , 18:65-68 (2000).  |
|                     |                  | Delaglio, "Adaptive Analysis and Multivariate Methods for Applications," <i>NMR Technologies: Development and Applications for Drug Discovery</i> , Sheraton Inner Harbor Hotel, Baltimore, Maryland (November 4-5, 1999).          |
|                     |                  | Detlefsen et al., "Molecular Flexibility Profiling Using NMR Spectroscopy," <i>Current Medicinal Chemistry</i> , 6(5):353-358 (1999).   |
|                     |                  | Fairbanks et al., "Purification and Structural Characterization of the CD11b/CD18 Integrin $\alpha$ Subunit I Domain Reveals a Folded Conformation in Solution," <i>FEBS Letters</i> , 369(2-3):197-201 (1995).                     |
|                     |                  | Farly et al., "SMASH: Small Molecule NMR Conference," Presentation Materials for SMASH'99, Argonne, IL (August 15-18, 1999).  |
| ✓                   |                  | Farmer II et al., "Localizing the NADP <sup>+</sup> Binding Site on the MurB Enzyme by NMR," <i>Nature Structural Biology</i> , 3(12):995-997 (1996).   |

|  |                        |
|--|------------------------|
| <b>EXAMINER</b>  | <b>Date Considered</b> |
| /Mark Shibuya/   | 06/21/2006             |
| *Examiner: Initial if citation considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant. |                        |

|   |  |                        |
|---|--|------------------------|
| <b>INFORMATION<br/>DISCLOSURE<br/>STATEMENT</b> | Atty. Docket No.: 6283.N DV1   | Serial No.: 10/694,385 |
|   | Applicant(s): Stockman et al.  | Confirmation No.: 5758 |
|   | Application Filing Date: Oct. 27, 2003   | Group: 1645            |
|   | Information Disclosure Statement<br>submitted (via facsimile) on: <b>MARCH 1, 2004</b> |                        |

| Examiner<br>Initial | Copy<br>Enclosed | Document Description   |
|---------------------|------------------|--|
| MS                  |                  | Fecik et al., "The Search for Orally Active Medications Through Combinatorial Chemistry," <i>Medicinal Research Reviews</i> , 1998;18:149-185.   |
|                     |                  | Fejzo et al., "The SHAPES Strategy: An NMR Based Approach for Lead Generation in Drug Discovery," <i>Chemistry and Biology</i> , 1999;6(10):755-769 and Abstract MIIA-4, <i>Proceedings of the 18<sup>th</sup> International Conference on Magnetic Resonance in Biological Systems</i> , Tokyo Metropolitan University, Tokyo, Japan, 3 pages (August 23-28, 1998). |
|                     |                  | Fesik, "NMR Structure-Based Drug Design," <i>Journal of Biomolecular NMR</i> , 3(3):261-269 (1993).  |
|                     |                  | Freeman et al., "Proton-Detected <sup>15</sup> N NMR Spectroscopy and Imaging," EPO abstract, XP 002029543, from <i>Journal of Magnetic Resonance, Series B</i> , 102(2):183-192, 1 page (1993).   |
|                     |                  | Freeman et al., "Proton-Detected <sup>15</sup> N NMR Spectroscopy and Imaging," <i>Journal of Magnetic Resonance, Series B</i> , 102(2):183-192 (1993).  |
|                     |                  | Ghose et al., "A Knowledge-Based Approach in Designing Combinatorial or Medicinal Chemistry Libraries for Drug Discovery. 1. A Qualitative and Quantitative Characterization of Known Drug Databases," <i>J. Comb. Chem.</i> , 1:55-68 (1999).   |
|                     |                  | Gonnella et al., "Isotope-Filtered Affinity NMR," <i>Journal of Magnetic Resonance</i> , 131:336-338 (1998).   |
|                     |                  | Gounarides et al., "Nuclear Magnetic Resonance Chromatography: Applications of Pulse Field Gradient Diffusion NMR to Mixture Analysis and Ligand-Receptor Interactions," <i>Journal of Chromatography B</i> , 725(1):79-90 (1999).   |
|                     |                  | Grzesiek et al., "The Importance of Not Saturating H <sub>2</sub> O in Protein NMR. Application to Sensitivity Enhancement and NOE Measurements," <i>Journal of the American Chemical Society</i> , 115(26):12593-12594 (1993).  |
| ↓                   |                  | Hajduk et al., "One Dimensional Relaxation- and Diffusion-Edited NMR Methods for Screening Compounds That Bind to Macromolecules," <i>J. Am. Chem. Soc.</i> , 119:12257-12261 (1997).  |

|  |  |
|--|--|
| <b>EXAMINER</b><br><br>/Mark Shibuya/  | <b>Date Considered</b><br><br>06/21/2006 |
| *Examiner: Initial if citation considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant. |  |

|   |   |                        |
|---|---|------------------------|
| <b>INFORMATION<br/>DISCLOSURE<br/>STATEMENT</b> | Atty. Docket No.: 6283.N DV1  | Serial No.: 10/694,385 |
|   | Applicant(s): Stockman et al.   | Confirmation No.: 5758 |
|   | Application Filing Date: Oct. 27, 2003  | Group: 1645            |
|   | Information Disclosure Statement<br>submitted (via facsimile) on: MARCH 1, 2004 |                        |

| Examiner<br>Initial | Copy<br>Enclosed | Document Description  |
|---------------------|------------------|---|
| MS                  |                  | Hajduk et al., "NMR-Based Discovery of Lead Inhibitors That Block DNA Binding of the Human Papillomavirus E2 Protein," <i>J. Med. Chem.</i> , 1997;40(20): 3144-3150.   |
|                     |                  | Hajduk et al., "High-Throughput Nuclear Magnetic Resonance-Based Screening," <i>J. Med. Chem.</i> , 42:2315-2317 (1999).  |
|                     |                  | Hajduk et al., "Integration of NMR and High-Throughput Screening," <i>Comb. Chem. High Throughput Screen</i> , Dec;5(8):613-621 (2002).   |
|                     |                  | Hegde et al., "Crystal Structure at 1.7 Å of the Bovine Papillomavirus-1 E2 DNA-Binding Domain Bound to its DNA Target," <i>Nature</i> , 359(6395):505-512 (1992).  |
|                     |                  | Hegde et al., "Crystal Structure of the E2 DNA-Binding Domain from Human Papillomavirus Type 16: Implications for Its DNA Binding-Site Selection Mechanism," <i>Journal of Molecular Biology</i> , 284(5):1479-1489 (1998). |
|                     |                  | Henrichsen et al., "Bioaffinity NMR Spectroscopy: Identification of an E-Selectin Antagonist in a Substance Mixture by Transfer NOE," <i>Angewandte Chemie, International Edition</i> , 38(1/2):98-102 (1999).              |
|                     |                  | Holmes et al., "Development of a Model for Classification of Toxin-Induced Lesions Using <sup>1</sup> H NMR Spectroscopy of Urine Combined with Pattern Recognition," <i>NMR in Biomedicine</i> , 11:235-244 (1998).        |
|                     |                  | Hwang et al., "Water Suppression That Works. Excitation Sculpting Using Arbitrary Waveforms and Pulsed Field Gradients," <i>J. Magn. Reson. A</i> 112:275-279 (1995).   |
|                     |                  | Kallen et al., "Structural Basis for LFA-1 Inhibition Upon Lovastatin Binding to the CD11a I-Domain," <i>J. Mol. Biol.</i> , 292:1-9 (1999).  |
|                     |                  | Kasukawa et al., "A Fifteen-Amino-Acid Peptide Inhibits Human Papillomavirus E1-E2 Interaction and Human Papillomavirus DNA Replication In Vitro," <i>Journal of Virology</i> , 72(10):8166-8173 (1998).                    |
| ✓                   |                  | Keifer, "High-Resolution NMR Techniques for Solid-Phase Synthesis and Combinatorial Chemistry," <i>Drug Discovery Today</i> , 2:468-478 (1997).   |

|  |                        |
|--|------------------------|
| <b>EXAMINER</b>  | <b>Date Considered</b> |
| /Mark Shibuya/   | 06/21/2006             |
| *Examiner: Initial if citation considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant. |                        |

|   |   |                        |
|---|---|------------------------|
| <b>INFORMATION<br/>DISCLOSURE<br/>STATEMENT</b> | Atty. Docket No.: 6283.N DV1  | Serial No.: 10/694,385 |
|   | Applicant(s): Stockman et al.   | Confirmation No.: 5758 |
|   | Application Filing Date: Oct. 27, 2003  | Group: 1645            |
|   | Information Disclosure Statement<br>submitted (via facsimile) on: <b>MARCH</b> <u>  </u> , 2004 |                        |

| Examiner<br>Initial | Copy<br>Enclosed | Document Description   |
|---------------------|------------------|--|
| MS                  |                  | Keifer, "New Methods for Obtaining High-Resolution NMR Spectra of Solid-Phase Synthesis Resins, Natural Products, and Solution-State Combinatorial Chemistry Libraries," <i>Drugs of the Future</i> , 23:301-317 (1998).                             |
|                     |                  | Keifer, "NMR Tools for Biotechnology," <i>Current Opinion in Biotechnology</i> , 10:34-41 (1999).  |
|                     |                  | Keifer et al., "Direct-Injection NMR (DI-NMR): A Flow NMR Technique for the Analysis of Combinatorial Chemistry Libraries," <i>J. Comb. Chem.</i> , 2:151-171 (2000).  |
|                     |                  | Lennon et al., "Hemoglobin Affinity for 2,3-Bisphosphoglycerate in Solutions and Intact Erythrocytes: Studies Using Pulsed-Field Gradient Nuclear Magnetic Resonance and Monte Carlo Simulations," <i>Biophysical Journal</i> , 67:2096-2109 (1994). |
|                     |                  | Liang et al., "Solution Structure of the DNA-Binding Domain of a Human Papillomavirus E2 Protein: Evidence for Flexible DNA-Binding Regions," <i>Biochemistry</i> , 35(7):2095-2103 (1996).  |
|                     |                  | Liepinsh et al., "Organic Solvents Identify Specific Ligand Binding Sites on Protein Surfaces," <i>Nature Biotechnology</i> , 15(3):264-268 (1997).  |
|                     |                  | Lin et al., "Diffusion-Edited NMR-Affinity NMR for Direct Observation of Molecular Interactions," <i>J. Am. Chem. Soc.</i> , 119:5249-5250 (1997).   |
|                     |                  | Lin et al., "Screening Mixtures by Affinity NMR," <i>J. Org. Chem.</i> , 62:8930-8931 (1997).  |
|                     |                  | Lindon et al., "Direct Coupling of Chromatographic Separations to NMR Spectroscopy," <i>Progress in Nuclear Magnetic Resonance Spectroscopy</i> , 29:1-49 (1996).  |
|                     |                  | Lindon et al., "Directly Coupled HPLC-NMR and Its Application to Drug Metabolism," <i>Drug Metabolism Reviews</i> , 29:705-746 (1997).   |
| ↓                   |                  | Lipinski et al., "Experimental and Computational Approaches to Estimate Solubility and Permeability in Drug Discovery and Development Settings," <i>Advanced Drug Delivery Reviews</i> , 23:3-25 (1997).   |

|  |  |
|--|--|
| <b>EXAMINER</b><br><br>/Mark Shibuya/  | <b>Date Considered</b><br><br>06/21/2006 |
| *Examiner: Initial if citation considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant. |  |

|   |   |                               |
|---|---|-------------------------------|
| <b>INFORMATION<br/>DISCLOSURE<br/>STATEMENT</b> | <b>Atty. Docket No.:</b> 6283.N DV1   | <b>Serial No.:</b> 10/694,385 |
|   | <b>Applicant(s):</b> Stockman et al.  | <b>Confirmation No.:</b> 5758 |
|   | <b>Application Filing Date:</b> Oct. 27, 2003   | <b>Group:</b> 1645            |
|   | <b>Information Disclosure Statement<br/>submitted (via facsimile) on:</b> MARCH <u>1</u> , 2004 |                               |

| Examiner<br>Initial | Copy<br>Enclosed | Document Description  |
|---------------------|------------------|---|
| MS                  |                  | Liu et al., "High-Resolution Diffusion and Relaxation Edited One- and Two-Dimensional <sup>1</sup> H NMR Spectroscopy of Biological Fluids," <i>Analytical Chemistry</i> , 68(19):3370-3376 (1996).   |
|                     |                  | Martin et al., "Beyond Mere Diversity: Tailoring Combinatorial Libraries for Drug Discovery," <i>J. Comb. Chem.</i> , 1:32-45 (1999).   |
|                     |                  | Mazel et al., "Genetic Characterization of Polypeptide Deformylase, a Distinctive Enzyme of Eubacterial Translation," <i>The EMBO Journal</i> , 13:914-923 (1994).  |
|                     |                  | McBride et al., "The Papillomavirus E2 Regulatory Proteins," <i>The Journal of Biological Chemistry</i> , 266(28):18411-18414 (1991).   |
|                     |                  | Melacini et al., "Band-Selective Editing of Exchange-Relay in Protein-Water NOE Experiments," <i>J. Biomol. NMR</i> , 13:67-71 (1999).  |
|                     |                  | Melacini et al., "Water-Macromolecule Interactions by NMR: a Quadrature-Free Constant-Time Approach and Its Application to C12," <i>J. Biomol. NMR</i> , 15:189-201 (1999).   |
|                     |                  | Meyer et al., "Screening Mixtures for Biological Activity by NMR," <i>Eur. J. Biochem.</i> , 246:705-709 (1997).  |
|                     |                  | Moore, "NMR Techniques for Characterization of Ligand Binding: Utility for Lead Generation and Optimization in Drug Discovery," <i>Biopolymer Peptide Science</i> , 51:221-243 (1999).  |
|                     |                  | Moore, "NMR Screening in Drug Discovery," <i>Current Opinion in Biotechnology</i> , 10(1):54-58 (1999).   |
|                     |                  | Morris et al., "Diffusion-Ordered Two-Dimensional Nuclear Magnetic Resonance Spectroscopy," <i>Journal of the American Chemical Society</i> , 114(8):3139-3141 (1992).  |
|                     |                  | Morris et al., "Resolution of Discrete and Continuous Molecular Size Distributions by Means of Diffusion-Ordered 2D NMR Spectroscopy," <i>Journal of the American Chemical Society</i> , 115(10):4291-4299 (1993).  |
| ✓                   |                  | Neri et al., " <sup>1</sup> H, <sup>13</sup> C and <sup>15</sup> N Backbone Assignments of Cyclophilin When Bound to Cyclosporin A (CsA) and Preliminary Structural Characterization of the CsA Binding Site," <i>FEBS Letters</i> , 294(1,2):81-88 (1991). |

|  |  |
|--|--|
| <b>EXAMINER</b><br><br>/Mark Shibuya/  | <b>Date Considered</b><br><br>06/21/2006 |
| *Examiner: Initial if citation considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant. |  |



|   |  |                               |
|---|--|-------------------------------|
| <b>INFORMATION<br/>DISCLOSURE<br/>STATEMENT</b> | <b>Atty. Docket No.:</b> 6283.N DV1  | <b>Serial No.:</b> 10/694,385 |
|   | <b>Applicant(s):</b> Stockman et al.   | <b>Confirmation No.:</b> 5758 |
|   | <b>Application Filing Date:</b> Oct. 27, 2003  | <b>Group:</b> 1645            |
|   | Information Disclosure Statement<br>submitted (via facsimile) on: <b>MARCH 1, 2004</b> |                               |

| Examiner<br>Initial | Copy<br>Enclosed | Document Description  |
|---------------------|------------------|---|
| MS                  |                  | Nicholson et al., "'Metabonomics': Understanding the Metabolic Responses of Living Systems to Pathophysiological Stimuli Via Multivariate Statistical Analysis of Biological NMR Spectroscopic Data," <i>Xenobiotica</i> , 29:1181-1189 (1999). |
|                     |                  | Otting et al., "Studies of Protein Hydration in Aqueous Solution by Direct NMR Observation of Individual Protein-Bound Water Molecules," <i>J. Am. Chem. Soc.</i> , 111:1871-1875 (1989).   |
|                     |                  | Otting, "NMR Studies of Water Bound Biological Molecules," <i>Progr. NMR Spectrosc.</i> , 31:259-285 (1997).  |
|                     |                  | Otting et al., "Protein Hydration in Aqueous Solution," <i>Science</i> , 254(5034): 974-980 (1991).   |
|                     |                  | Pearlman et al., "Novel Software Tools for Chemical Diversity," <i>Perspectives in Drug Discovery and Design</i> , 09/10/11:339-353 (1998).   |
|                     |                  | Phelps et al., "Molecular Targets for Human Papillomaviruses: Prospects for Antiviral Therapy," <i>Antiviral Chemistry &amp; Chemotherapy</i> , 9(5):359-377 (1998).  |
|                     |                  | Piotto et al., "Gradient-Tailored Excitation for Single-Quantum NMR Spectroscopy of Aqueous Solutions," <i>Journal of Biomolecular NMR</i> , 2(6):661-665 (1992).   |
|                     |                  | Ponstingl et al., "Detection of Protein-Ligand NOEs With Small, Weakly Binding Ligands by Combined Relaxation and Diffusion Filtering," <i>Journal of Biomolecular NMR</i> , 9:441-444 (1997).  |
|                     |                  | Price, "Water Signal Suppression in NMR Spectroscopy," <i>Annual Reports on NMR Spectroscopy</i> (Ed., Webb, A.), Academic Press, New York, vol. 38, pp. 289-354 (1999).  |
|                     |                  | Rabenstein et al., "A Pulse Sequence for the Measurement of Spin-Lattice Relaxation Times of Small Molecules in Protein Solutions," <i>Journal of Magnetic Resonance</i> , 34:669-674 (1979).   |
| ↓                   |                  | Ross et al., "Fast-HMQC Using Ernst Angle Pulses: An Efficient Tool for Screening of Ligand Binding to Target Proteins," <i>Journal of Biomolecular NMR</i> , 10:389-396 (1997).  |

|   |  |
|---|--|
| <b>EXAMINER</b><br><br>/Mark Shibuya/   | <b>Date Considered</b><br><br>06/21/2006 |
| <small>*Examiner: Initial if citation considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.</small> |  |

|   |   |                        |
|---|---|------------------------|
| <b>INFORMATION<br/>DISCLOSURE<br/>STATEMENT</b> | Atty. Docket No.: 6283.N DV1  | Serial No.: 10/694,385 |
|   | Applicant(s): Stockman et al.   | Confirmation No.: 5758 |
|   | Application Filing Date: Oct. 27, 2003  | Group: 1645            |
|   | Information Disclosure Statement<br>submitted (via facsimile) on: MARCH 1, 2004 |                        |

| Examiner<br>Initial | Copy<br>Enclosed | Document Description   |
|---------------------|------------------|--|
| MS                  |                  | Ross et al., "Automation of NMR Measurements and Data Evaluation for Systematically Screening Interactions of Small Molecules with Target Proteins," <i>Journal of Biomolecular NMR</i> , 16:139-146 (2000).   |
|                     |                  | Sadowski et al., "A Scoring Scheme for Discriminating Between Drugs and Nondrugs," <i>J. Med. Chem.</i> , 41:3325-3329 (1998).   |
|                     |                  | Scherf et al., "A T <sub>1</sub> -Filtered Two-Dimensional Transferred NOE Spectrum for Studying Antibody Interactions with Peptide Antigens," <i>Biophysical Journal</i> , 64:754-761 (1993).   |
|                     |                  | Shapiro et al., "High Resolution NMR for Screening Ligand/Protein Binding," <i>Current Opinion in Drug Discovery &amp; Development</i> , 2:396-400 (1999).   |
|                     |                  | Shuker, "Discovering High-Affinity Ligands for Proteins: SAR by NMR," <i>Science</i> , 274:1531-1534 (1996).   |
|                     |                  | Sobol et al., "Solvent Magnetization Artifacts in High-Field NMR Studies of Macromolecular Hydration," <i>J. Magn. Reson.</i> 1998 Feb;130(2):262-271.   |
|                     |                  | Spraul et al., "High-Throughput Flow-Injection NMR and Its Applications," <i>Bruker Report</i> (1999).   |
|                     |                  | Spraul et al., "Flow Injection Proton Nuclear Magnetic Resonance Spectroscopy Combined With Pattern Recognition Methods: Implications for Rapid Structural Studies and High Throughput Biochemical Screening," <i>Analytical Communications</i> , 34:339-341 (1997).         |
|                     |                  | Stilbs, "Molecular Self-Diffusion Coefficients in Fourier Transform Nuclear Magnetic Resonance Spectrometric Analysis of Complex Mixtures," <i>Analytical Chemistry</i> , 53(13):2135-2137 (1981).   |
|                     |                  | Stockman et al., " <sup>1</sup> H and <sup>15</sup> N Resonance Assignments and Solution Secondary Structure of Oxidized <i>Desulfovibrio vulgaris</i> Flavodoxin Determined by Heteronuclear Three-Dimensional NMR Spectroscopy," <i>J. Biomol. NMR</i> , 3:133-149 (1993). |
| ✓                   |                  | Stockman, "NMR Spectroscopy as a Tool for Structure-Based Drug Design," <i>Progress in Nuclear Magnetic Resonance Spectroscopy</i> , 33:109-151 (1998).  |

|  |                        |
|--|------------------------|
| <b>EXAMINER</b>  | <b>Date Considered</b> |
| /Mark Shibuya/   | 06/21/2006             |
| *Examiner: Initial if citation considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant. |                        |

|   |  |                        |
|---|--|------------------------|
| <b>INFORMATION<br/>DISCLOSURE<br/>STATEMENT</b> | Atty. Docket No.: 6283.N DV1   | Serial No.: 10/694,385 |
|   | Applicant(s): Stockman et al.  | Confirmation No.: 5758 |
|   | Application Filing Date: Oct. 27, 2003   | Group: 1645            |
|   | Information Disclosure Statement<br>submitted (via facsimile) on: <b>MARCH 1, 2004</b> |                        |

| Examiner<br>Initial | Copy<br>Enclosed | Document Description   |
|---------------------|------------------|--|
| MS                  |                  | Stockman, "Applications of Flow NMR Spectroscopy to Monitor Binding of Small Molecules to Proteins," Innovative Computational Applications: The Interface of Library Design, Bioinformatics, Structure Based Drug Design and Virtual Screening, Biotechnology Division, Institute for International Research, San Francisco, CA, Oct. 25-27, 1999. |
|                     |                  | Stockman, "Applications of flow NMR Spectroscopy to Monitor Binding of Small Molecules to Proteins," NMR Technologies: Development and Applications for Drug Discovery, Cambridge Healthtech Institute's Second International, Baltimore, MD, Nov. 4-5, 1999.  |
|                     |                  | Stockman, "Flow NMR Spectroscopy in Drug Discovery," <i>Current Opinion in Drug Discovery &amp; Development</i> , 3:269-274 (2000).  |
|                     |                  | Stockman, "Applications of Flow NMR Spectroscopy to Monitor Binding of Small Molecules to Proteins," NMR in the Drug Discovery Pipeline, IBC, London, UK, May 8-9, 2000.   |
|                     |                  | Stockman et al., "Screening of Compound Libraries for Protein Binding Using Flow-Injection Nuclear Magnetic Resonance Spectroscopy," <i>Methods Enzymol.</i> 2001;338:230-246.   |
|                     |                  | Tatusova et al., "BLAST 2 Sequences, a New Tool for Comparing Protein and Nucleotide Sequences," <i>FEMS Microbiology Letters</i> , 174(2):247-250 (1999).   |
|                     |                  | Teague et al., "The Design of Leadlike Combinatorial Libraries," <i>Angew. Chem. Int. Ed.</i> 38:3743-3748 (1999).   |
|                     |                  | Veeraraghavan et al., " <sup>1</sup> H, <sup>15</sup> N, and <sup>13</sup> C NMR Resonance Assignments for the DNA-Binding Domain of the BPV-1 E2 Protein," <i>Journal of Biomolecular NMR</i> , 11(4):457-458 (1998).   |
|                     |                  | Veeraraghavan et al., "Structural Correlates for Enhanced Stability in the E2 DNA-Binding Domain from Bovine Papillomavirus," <i>Biochemistry</i> , 1999; 38(49):16115-16124.  |
| ✓                   |                  | Vogler et al., "Rapid Communications," <i>Journal of Natural Products</i> , 61:175-178 (1998).   |

|  |  |
|--|--|
| <b>EXAMINER</b><br><br>/Mark Shibuya/  | <b>Date Considered</b><br><br>06/21/2006 |
| *Examiner: Initial if citation considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant. |  |

|   |  |                        |
|---|--|------------------------|
| <b>INFORMATION<br/>DISCLOSURE<br/>STATEMENT</b> | Atty. Docket No.: 6283.N DVI   | Serial No.: 10/694,385 |
|   | Applicant(s): Stockman et al.  | Confirmation No.: 5758 |
|   | Application Filing Date: Oct. 27, 2003   | Group: 1645            |
|   | Information Disclosure Statement<br>submitted (via facsimile) on: <b>MARCH 1, 2004</b> |                        |

| Examiner<br>Initial | Copy<br>Enclosed | Document Description  |
|---------------------|------------------|---|
| MS                  |                  | Wang et al., "Solution Studies of Staphylococcal Nuclease H124L. 2. <sup>1</sup> H, <sup>13</sup> C, and <sup>15</sup> N Chemical Shift Assignments for the Unligated Enzyme and Analysis of Chemical Shift Changes that Accompany Formation of the Nuclease-Thymidine 3', 5'-Bisphosphate-Calcium Ternary Complex <sup>1,2</sup> ," <i>Biochemistry</i> , 31:921-936 (1992). |
|                     |                  | Wang et al., "Toward Designing Drug-Like Libraries: A Novel Computational Approach for Prediction of Drug Feasibility of Compounds," <i>J. Comb. Chem.</i> , 1: 524-533 (1999).   |
|                     |                  | Warr, "Combinatorial Chemistry and Molecular Diversity. An Overview," <i>J. Chem. Inf. Comput. Sci.</i> , 1997;37:134-140.  |
|                     |                  | Watanabe et al., "Direct-Coupling of FT-NMR to High Performance Liquid Chromatography," <i>Proc. Japan Acad. Ser B</i> , 54:194-199 (1978).   |
|                     |                  | Watt et al., "Comparison of the Crystal Structures of a Flavodoxin in its Three Oxidation States at Cryogenic Temperatures," <i>J. Mol. Biol.</i> , 218:195-208 (1991).   |
|                     |                  | Wider et al., "Proton-Proton Overhauser Effects of Receptor-Bound Cyclosporin A Observed With the Use of a Heteronuclear-Resolved Half-Filter Experiment," EPO abstract, XP 002029543, from <i>Journal of the American Chemical Society</i> , 113(12):4676-4678, 2 pages (1991).  |
|                     |                  | Wider et al., "Proton-Proton Overhauser Effects of Receptor-Bound Cyclosporin A Observed with the Use of a Heteronuclear-Resolved Half-Filter Experiment," <i>Journal of the American Chemical Society</i> , 113(12):4676-4678 (1991).  |
|                     |                  | Wider, "Structure Determination of Biological Macromolecules in Solution Using Nuclear Magnetic Resonance Spectroscopy," <i>BioTechniques</i> , 29(6):1278-1294 (2000).   |
|                     |                  | Williams, Abstracts of Papers, Part 1, 218 <sup>th</sup> ACS National Meeting at New Orleans, LA (August 22-26, 1999).  |
|                     |                  | Wishart et al., "The <sup>13</sup> C Chemical-Shift Index: A Simple Method for the Identification of Protein Secondary Structure Using <sup>13</sup> C Chemical-Shift Data," <i>Journal of Biomolecular NMR</i> , 4(2):171-180 (1994).  |
| ✓                   |                  | Wishart et al., "Protein Chemical Shift Analysis: A Practical Guide," <i>Biochemistry and Cell Biology</i> , 76(2/3):153-163 (1998).  |

EXAMINER

/Mark Shibuya/

Date Considered

06/21/2006

\*Examiner: Initial if citation considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

|   |   |                        |
|---|---|------------------------|
| <b>INFORMATION<br/>DISCLOSURE<br/>STATEMENT</b> | Atty. Docket No.: 6283.N DV1  | Serial No.: 10/694,385 |
|   | Applicant(s): Stockman et al.   | Confirmation No.: 5758 |
|   | Application Filing Date: Oct. 27, 2003  | Group: 1645            |
|   | Information Disclosure Statement<br>submitted (via facsimile) on: MARCH 1, 2004 |                        |

| Examiner<br>Initial | Copy<br>Enclosed | Document Description  |
|---------------------|------------------|---|
| MS                  |                  | Wolfender et al., "LC/NMR in Natural Products Chemistry," <i>Current Organic Chemistry</i> , 2:575-596 (1998).  |
| MS                  |                  | Wu et al., "An Improved Diffusion-Ordered Spectroscopy Experiment Incorporating Bipolar-Gradient Pulses," <i>J. Magn. Reson. Ser. A</i> , 1995;115:260-264. |
|                     |                  |   |
|                     |                  |   |
|                     |                  |   |

|   |  |
|---|--|
| <b>EXAMINER</b><br><br>/Mark Shibuya/   | <b>Date Considered</b><br><br>06/21/2006 |
| <small>*Examiner: Initial if citation considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.</small> |  |